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# Self-monitoring as a familial vulnerability marker for psychosis: an analysis of patients, unaffected siblings and healthy controls

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**Background.** Alterations in self-monitoring have been reported in patients with psychotic disorders, but it remains unclear to what degree they represent true indicators of familial vulnerability for psychosis.

**Method.** An error-correction action-monitoring task was used to examine self-monitoring in 42 patients with schizophrenia, 32 of their unaffected siblings and 41 healthy controls.

**Results.** Significant between-group differences in self-monitoring accuracy were found ( $\chi^2=29.3$ ,  $p<0.0001$ ), patients performing worst and unaffected siblings performing at an intermediate level compared to controls (all between-group differences  $p<0.05$ ). In the combined group of healthy controls and unaffected siblings, detection accuracy was associated with positive schizotypy as measured by the Structured Interview for Schizotypy – Revised (SIS-R) ( $\beta=-0.16$ ,  $\text{s.e.}=0.07$ ,  $p=0.026$ ), but not with negative schizotypy ( $\beta=-0.05$ ,  $\text{s.e.}=0.12$ ,  $p=0.694$ ). In patients, psychotic symptoms were not robustly associated with detection accuracy ( $\beta=-0.01$ ,  $\text{s.e.}=0.01$ ,  $p=0.094$ ), although stratified analysis revealed suggestive evidence for association in patients not currently using antipsychotic medication ( $\beta=-0.03$ ,  $\text{s.e.}=0.01$ ,  $p=0.052$ ), whereas no association was found in patients on antipsychotic medication ( $\beta=-0.01$ ,  $\text{s.e.}=0.01$ ,  $p=0.426$ ). A similar pattern of associations was found for negative symptoms.

**Conclusions.** Alterations in self-monitoring may be associated with familial risk and expression of psychosis. The association between psychotic symptoms and self-monitoring in patients may be affected by antipsychotic medication, which may explain previous inconsistencies in the literature.

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**Key words:** Psychosis, schizophrenia, self-monitoring.

## Introduction

Self-monitoring is a specific type of source monitoring (Johnson *et al.* 1993; Brebion *et al.* 2007; Versmissen *et al.* 2007b) that enables the person to distinguish self-generated actions from those elicited by external stimuli (Frith, 1987; Knoblich *et al.* 2004). It has been proposed that the internal monitoring of speech generation and inner thoughts is analogous to the monitoring of motor actions (Frith, 1987, 1992, 1996; Blakemore *et al.* 2000; Seal *et al.* 2004; Fu *et al.* 2006; Jones & Fernyhough, 2007). According to this model,

‘thinking’ is operationalized as an action with a clear intention, thus providing a sense of agency for thoughts (Frith, 1992; Frith *et al.* 1998, 2000; Gallagher, 2004). Misattributions of self-generated thoughts as voices coming from the outside could provide a plausible mechanism underlying the hallucinatory process in psychosis (Shergill *et al.* 2000). A deficit in the process of self-monitoring thoughts may therefore result in thought insertion and auditory hallucinations (Frith, 1992; Brebion *et al.* 1998, 2005; Ditman & Kuperberg, 2005). Several studies in patients with schizophrenia have indeed provided evidence that a deficit in self-monitoring of one’s own cognitive actions is associated with psychotic symptoms e.g. (Harvey, 1985; Bentall *et al.* 1991; Daprati *et al.* 1997; Blakemore *et al.* 2000; Keefe *et al.* 2002; Brunelin *et al.* 2006; Johns *et al.* 2006; Costafreda *et al.* 2008), although

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not all studies were able to find an association between psychotic symptoms and alterations in self-monitoring (Vinogradov *et al.* 1997; Fournier *et al.* 2001; Li *et al.* 2002; Versmissen *et al.* 2007a).

Importantly, studies suggesting that alterations in self-monitoring are associated with psychotic disorder were unable to consistently address the question of whether self-monitoring alterations are a vulnerability marker for psychotic disorder or merely a state marker of acute psychotic symptoms. Johns *et al.* (2006) found a self-monitoring deficit in patients currently experiencing hallucinations but not in patients who previously experienced hallucinations. Franck *et al.* (2000) found worse detection accuracy in patients experiencing hallucinations during the experiment compared to non-hallucinating patients. Several studies have used functional neuroimaging paradigms to detect abnormal regional activation in hallucination-prone patients when performing tasks engaging the verbal self-monitoring system, compared to non-hallucination prone patients with schizophrenia and healthy controls, focusing on various regions such as the Broca area, the anterior cingulate, the left temporal cortex and the supplementary motor area (SMA) (McGuire *et al.* 1995, 1996; Schnell *et al.* 2008). These studies reported both under- and overactivation associated with self-monitoring alterations (Shergill *et al.* 2000; Schnell *et al.* 2008; Raij *et al.* 2009), and these findings were hypothesized to result from a pathological increase in microstructural elements supporting excitatory neurotransmission, leading to instability (for a detailed discussion, see Bates *et al.* 2002).

Some studies in healthy control subjects have found evidence suggesting that alterations in self-monitoring may be associated with vulnerability for psychosis rather than acute psychosis *per se*. For example, several studies have found that healthy individuals who experienced hallucinations displayed more reality or action-monitoring errors compared to normal subjects who had not experienced hallucinations (Bentall & Slade, 1985; Rankin & O'Carroll, 1995; Laroie *et al.* 2004; Debbané *et al.* 2009). An additional argument that alterations in self-monitoring are not merely epiphenomena of acute psychosis is that some studies have also reported an inverse association between self-monitoring and negative symptoms (Brebion *et al.* 1999, 2002, 2005). Negative symptoms are thought to be stable over time (Pfohl & Winokur, 1982; Pogue-Geile & Harrow, 1985; Katsanis *et al.* 1990; Keefe *et al.* 1991; Rey *et al.* 1994; Dollfus & Petit, 1995) and to index genetic liability to psychotic disorder better than positive symptoms (Tsuang *et al.* 1991; Fanous *et al.* 2001; Schurhoff *et al.* 2003).

Further evidence suggesting that self-monitoring alterations are associated with vulnerability for

psychosis may be derived from studies investigating groups at risk for psychotic disorder such as first-degree relatives of patients with schizophrenia (Johns & van Os, 2001; Hanssen *et al.* 2006) or individuals with psychometric risk states. To our knowledge, only two studies have used this approach. Johns *et al.* (2009) found a significant difference in self-monitoring in persons with an at-risk mental state (ARMS) compared to healthy controls, and Versmissen *et al.* (2007b) found evidence for self-monitoring alterations in three psychometrically and genetically defined at-risk groups compared to controls.

Of note is that the use of verbal recognition or signal detection tasks, as applied in previous research (Bentall & Slade, 1985; Rankin & O'Carroll, 1995; Morrison & Haddock, 1997; Brebion *et al.* 1998, 2000, 2005; Ragland *et al.* 2003; Heinrichs & Vaz, 2004; Johns *et al.* 2009), may yield biased results because performance on these tasks is strongly associated with cognitive performance in general and could thus be influenced by characteristics such as verbal intelligence, time elapsed between acquisition and recognition, memory dysfunction and executive dysfunction (Johnson *et al.* 1993; Seal *et al.* 1997). Therefore, in the present study we used an action-monitoring task, which may be less biased by cognitive performance, in patients with schizophrenia, their unaffected siblings and healthy controls. We hypothesized that: (1) patients with a non-affective psychotic disorder would show worse self-monitoring compared to healthy controls; (2) the unaffected siblings would show intermediate accuracy, supporting the notion of self-monitoring alterations as a vulnerability marker for psychosis; (3) self-monitoring accuracy would be inversely related to psychotic symptoms at the level of subclinical positive schizotypy in unaffected groups as well as at the level of psychotic symptoms in the patients; and (4) self-monitoring alterations would also be associated with negative symptoms in both patients and in non-affected groups.

Furthermore, it was hypothesized that patients do not automatically change their own movements to compensate for the action change by the computer compared to healthy controls or their first-degree relatives, as reported consistently in previous research (for an overview, see Jeannerod, 2009).

## Method

### Sample

The study sample consisted of 42 patients with a non-affective psychotic disorder according to DSM-IV-TR, 32 unaffected siblings of the patients and 49 control

subjects. Written informed consent conforming to the local ethics committee guidelines was obtained from all subjects. Inclusion criteria were: fluency in Dutch; aged 16–55 years; and, for patients, first contact with mental health facilities within the past 10 years. For the controls, the occurrence of a psychotic disorder in either the control or a first-degree family member was considered an exclusion criterion.

Patients were recruited through the Community Mental Health Centres and the Psychiatric Hospitals of the catchment area (South Limburg, The Netherlands). All unaffected siblings were sampled through participating patients. Twenty-two families contributed one unaffected sibling and one patient, two families contributed two siblings and one patient, one family contributed two patients, seven unaffected siblings participated without their patient sibling and 16 patients participated without an unaffected sibling. Control subjects were recruited through random mailings in nearby municipalities and through advertisements.

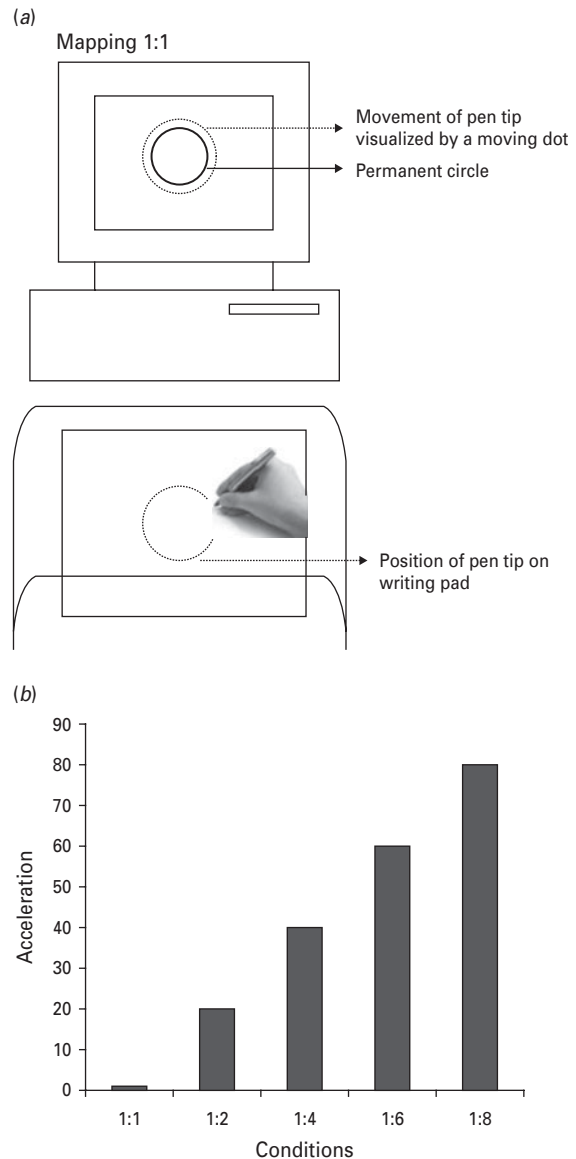
The 42 patients had DSM-IV-TR diagnoses of schizophrenia ( $n=32$ ), schizo-affective disorder ( $n=4$ ), brief psychotic disorder ( $n=5$ ) and psychotic disorder not otherwise specified (NOS) ( $n=1$ ). Five siblings and five healthy controls had had one episode of major depressive disorder (MDD), in full remission at the time of testing, and two controls had had one episode of MDD, in partial remission at the time of testing. The remaining siblings and controls had no DSM-IV diagnosis.

### Procedures

To study the hypothesis that symptoms are associated with problems in the central monitoring of action (Frith & Done, 1989), studies on error correction can be used to test the source-monitoring system. These studies test the central monitoring system because the error correction is too rapid to use exteroceptive feedback. In this study, an error correction task was used as described previously (Knoblich & Kircher, 2004; Knoblich *et al.* 2004).

Seated in front of a computer screen, at a distance of about 60 cm, subjects were exposed to the image of a full circle (see Fig. 1a). A covered writing pad was located between the subject and the computer screen. By covering the writing pad we could ensure that the subjects could not observe their own hand movements while they were drawing with a pen.

The computer screen was an Apple Vision (Apple, USA) 17-inch monitor with a horizontal resolution of 800 pixels and a vertical resolution of 600 pixels. The vertical sync frequency was 75 Hz. The movement of the pen tip was recorded using a pressure-sensitive



**Fig. 1.** (a) Visual representation of the task. Each separate trial consisted of drawing five circles around the full, permanent circle that remained on the screen during the whole trial. Participants were instructed to interrupt the test by lifting their pen from the drawing map as soon as they noticed a difference between the actual movement they made on the drawing map and the representation they saw on the computer screen. (b) The different conditions used in the error correction task. In the interval between the sixth and the eighth second, the movement of the dot on the screen was accelerated relative to the movement of the pen tip on the drawing map by 20, 40, 60 or 80%.

Wacom writing pad (Wacom Europe GmbH, Germany) with a sampling rate of 75 Hz, a horizontal resolution of 15000 dots and a vertical resolution of 11250 dots. An Apple Power PC controlled these devices. The sampling rate of the writing pad was

synchronized with the screen refresh rate. Hence, the constant delay between the visual effect and the movement of the pen tip was about 13 ms.

The experiment was made up of four tasks, which were presented in the following order.

#### *Tracking task*

The objective of the first task (20 trials) was to assess the subject's tracking performance. In each trial the subject tracked a circular target, which moved with constant velocity, using the pen and the writing pad. The location of the pen tip was indicated by a solid circular dot. Neither the circular target nor the dot left a trace on the screen. In this task the mapping between the computer screen and the writing pad was 1:1, that is a similarity of 100% between the drawing made by the subject and the visual consequence shown on the computer screen. In each trial the circular target completed five circles around the full, permanent circle, which remained on the screen during the whole trial, with a velocity of 2 s per circle and an eccentricity of 9° visually for the full circle. Crossing the 12 o'clock position of the target was indicated by a short beep (200 ms, 1000 Hz).

#### *Training task*

In this task 10 trials with mapping changes (from 1:1 to 1:2) were introduced to the subjects, similar to the main experiment.

#### *Colour change detection task*

In this condition we could assess whether there were difficulties lifting the pen during a movement. Therefore, the subject had to lift the pen as rapidly as possible when the solid circular dot changed colour. This colour change occurred while in the fourth circle.

#### *Main experiment*

Each trial (120 in total) of the experiment started by successfully crossing a small quadratic box, located above the 12 o'clock position of the circle, accompanied by a short beep. Each separate trial was composed of drawing five circles around the full, permanent circle that remained on the screen during the whole trial. Whenever the dot that represented the pen tip passed the 12 o'clock point (after 2, 4, 6, 8 and 10 s), the same beep was heard.

Participants were instructed to interrupt the test by lifting their pen from the drawing map as soon as they noticed a difference between the actual movement they made on the drawing map and the representation they saw on the computer screen. A trial could come to an end by the lifting of the pen or automatically after 11 s if no pen lift occurred.

The position of the pen tip's coordinates were recorded at the time of each beep to determine to what extent the mapping change was compensated for in the movement. This position reflects the distance between the centre of the drawn circle and the pen position on the writing pad. Therefore, the position of the pen tip at each beep represents the radius of the circle the subject was ending at the time of the beep.

Five different conditions (20% of the trials each) were possible (see Fig. 1*b*). The first 6 s, the mapping between the writing pad and the screen was 1:1. In the interval between the sixth and the eighth second, the movement of the dot on the screen was accelerated relative to the movement of the pen tip on the drawing map by 20, 40, 60 or 80%. In 20% of the trials, no acceleration occurred.

#### *Instruments*

For all participating patients, the Operational Criteria Checklist for Psychotic Illness (OCCPI; McGuffin *et al.* 1991) was completed, based on case-note material, and also the Positive and Negative Syndrome Scale (PANSS) interview (Kay *et al.* 1987). Where necessary, additional information was derived from ward staff or case managers. Using the information in the OCCPI, the computerized program OPCRIT (McGuffin *et al.* 1991) yielded DSM-IV diagnoses.

The Structured Interview for Schizotypy – Revised (SIS-R) was used to measure the positive, negative and disorganization dimensions of the subclinical psychosis or 'schizotypy' phenotype (Vollema & Ormel, 2000; Pfeifer *et al.* 2009). Items are scored on a four-point scale ranging from absent (score 0) to severe (score 3). Positive schizotypy covers the items referential thinking, magical ideation, illusions and suspiciousness. Negative schizotypy contains the items social isolation, social anxiety, introversion, restricted affect, referential thinking and suspiciousness. Disorganization schizotypy encompasses the items goal directness of thinking, loosening of associations, and oddness.

Finally, two verbal subtests, Information and Arithmetic, and two performance subtests, Block Design and Symbol Search, of the Wechsler Adult Intelligence Scale III (WAIS-III; Wechsler, 1997) were used to obtain a measure of intelligence quotient (IQ; Blyler *et al.* 2000).

#### *Analysis*

##### *Associations between psychosis risk and self-monitoring ability*

A three-level ordinal group variable was constructed reflecting the risk for psychosis, with a value of 2 for

patients, 1 for relatives and 0 for controls. The number of correct pen lifts in the five conditions, representing a certain extent of mapping change (hereafter 'detection accuracy'), were saved as average values over all trials for each subject. Pen lifts in the baseline conditions are errors in self-monitoring (no transformation occurs), whereas pen lifts in the other conditions are indicative of adequate self-monitoring. Therefore, to create a measure of the individual's overall detection accuracy, four observations were created for each subject, reflecting the four velocity transformation conditions where transformation occurred (1:2, 1:4, 1:6, 1:8). To account for hierarchical clustering of detection accuracies across different conditions (1:2, 1:4, 1:6, 1:8), nested in individual participants, between-group differences in the pattern of detection accuracy were modelled using multilevel linear regression analysis using the XTREG routine in STATA version 11.0 (StataCorp, 2008), with detection accuracy as the dependent variable and 'group' and 'condition', and also their interaction term, as the independent variables. This analysis investigates the increase in detection accuracy with increasing acceleration (i.e. over the different conditions), as indicated by the 'group  $\times$  condition interaction'. This approach allows for the combined analysis of all self-monitoring data, which is a better solution than analysing the conditions separately. Age, sex and IQ were included as possible *a priori* confounders. To determine whether differences between individual groups were statistically significant, *post-hoc* Wald tests based on the group  $\times$  condition interaction model were used. Effect sizes were obtained by examining the appropriate linear combinations using the LINCOM routine.

#### *Associations between psychotic symptoms and self-monitoring ability*

To examine the hypothesis that higher levels of positive and negative symptoms are associated with worse action-monitoring in patients, multilevel linear regression analysis was applied, with detection accuracy as the dependent variable and psychotic symptoms, condition and their interaction term as independent variables. A similar analysis was conducted with SIS-R subclinical symptoms in controls and unaffected siblings. For this analysis, the controls and the unaffected siblings were combined to increase the sample size and variability, resulting in greater statistical power.

Given the known overlap between positive and negative symptoms, the multilevel regression analysis investigating the association between positive symptoms and self-monitoring was controlled for the presence of negative symptoms, as in a previous study (Brebion *et al.* 1999), and *vice versa*. In addition, age, sex

**Table 1.** Demographics

|                     | Controls | First-degree relatives | Patients | <i>p</i> value |
|---------------------|----------|------------------------|----------|----------------|
| No. of participants | 49       | 32                     | 42       |                |
| Male (%)            | 32.7     | 36.4                   | 66.7     | 0.003          |
| Age (years)         | 34.81    | 30.84                  | 33.48    | 0.293          |
| IQ                  | 114.18   | 100.59                 | 95.63    | <0.001         |

and IQ were included as possible *a priori* confounders. The role of antipsychotic use was also considered as potentially relevant because antipsychotics exert a direct effect on symptomatology and it may be that they also influence the ability to monitor one's motor actions given the effects on nigrostriatal dopaminergic neurotransmission. Therefore, the analysis in patients was additionally controlled for use of antipsychotic medication (yes/no) and subanalyses also investigated the association between detection accuracy and symptomatology in patients stratified for current use of antipsychotic medication. Two-way interactions between condition and symptoms stratified for antipsychotic use were derived from the three-way condition  $\times$  symptoms  $\times$  antipsychotic use interaction model.

#### *Compensation of mapping change*

The radial components, reflecting the radius of the pen tip at the time of each beep, were transcribed for each beep (five beeps per trial), with separate values for the various mapping change conditions. Because the velocity transformation occurred after radius 3, the compensation movement per velocity transformation was calculated by subtracting the mean radial component before the velocity transformation from the mean radial component after the velocity transformation. Similar to the detection accuracy, the mean compensation movements per velocity transformation per subject were reshaped into five observations per subject based on the latter five velocity transformation conditions. Between-group differences were assessed, with 'mean compensation' as the dependent variable and 'group' and 'condition' and also their interaction term as the independent variables.

## **Results**

### *Sample characteristics*

The error-correction task was carried out by 124 subjects: 42 patients with psychosis, 32 siblings and 49 healthy controls. Significant between-group differences were found for intelligence and sex (see Table 1).

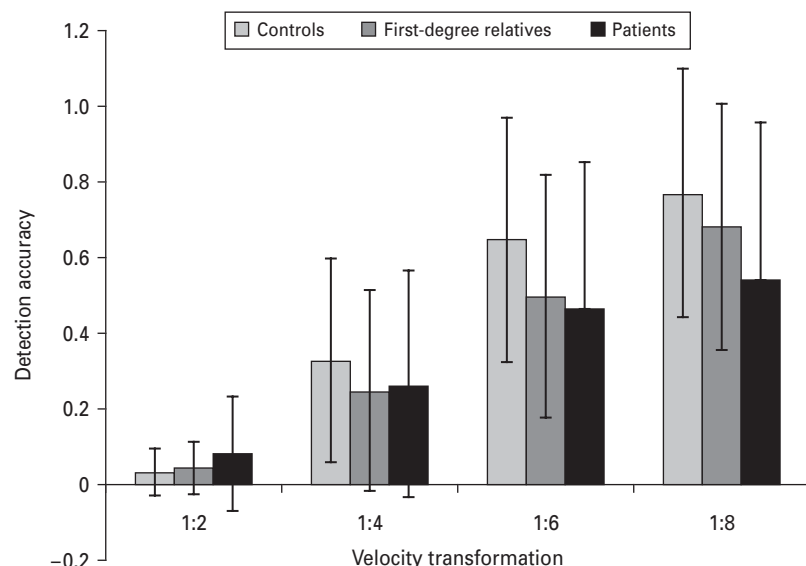


Fig. 2. Patterns of detection accuracy in different conditions across groups.

Twenty-nine patients with psychosis were currently using antipsychotic medication, 10 patients were not, and no information on antipsychotic use was available for three patients. There were no large or significant differences between patients with or without current antipsychotic treatment in positive ( $t = -0.28$ ,  $p = 0.778$ ) or negative symptoms ( $t = -0.90$ ,  $p = 0.374$ ).

### Self-monitoring ability and psychosis

The detection accuracy increased significantly in all groups as the transformations became larger ( $p < 0.001$  for all). Performance on the action-monitoring task was modestly associated with IQ in the patient group ( $\beta = 0.02$ ,  $\text{s.e.} = 0.01$ ,  $p = 0.033$ ) but not in the siblings ( $\beta = 0.004$ ,  $\text{s.e.} = 0.01$ ,  $p = 0.61$ ) or healthy controls ( $\beta = 0.01$ ,  $\text{s.e.} = 0.01$ ,  $p = 0.14$ ). Significant differences between groups were found in detection accuracy over the different conditions when covarying for age, sex and IQ [group  $\times$  condition interaction  $\chi^2(2) = 29.3$ ,  $p < 0.0001$ ], as depicted in Fig. 2 and Table 2. Patients showed the worst detection accuracy ( $\beta = 0.41$ ,  $\text{s.e.} = 0.04$ ), controls had the best ( $\beta = 0.68$ ,  $\text{s.e.} = 0.03$ ) and the siblings performed at a level that was intermediate between the other groups ( $\beta = 0.56$ ,  $\text{s.e.} = 0.04$ ). Differences between controls and patients were statistically significant [ $\chi^2(1) = 29.8$ ,  $p < 0.0001$ ], as were differences between controls and siblings [ $\chi^2(1) = 5.8$ ,  $p = 0.016$ ] and patients and siblings [ $\chi^2(1) = 7.0$ ,  $p = 0.008$ ]. Because psychomotor slowing, as measured by the Symbol-Digit Substitution task, was observed in both patients ( $\beta = -18.0$ ,  $\text{s.e.} = 1.57$ ,  $p < 0.01$ ) and unaffected siblings compared to the controls ( $\beta = -4.26$ ,  $\text{s.e.} = 1.69$ ,  $p = 0.012$ ), a sensitivity

analysis investigated action monitoring while covarying for psychomotor performance (in addition to the *a priori* confounders age, sex and IQ). This did not alter the results, indicating that the observed associations are not due to psychomotor slowing in the siblings and patients. Patients did not perceive changes significantly more often than their unaffected siblings or the controls in the 1:1 condition in which there was no acceleration (siblings:  $\beta = 0.02$ ,  $\text{s.e.} = 0.02$ ,  $p = 0.30$ ; controls:  $\beta = 0.02$ ,  $\text{s.e.} = 0.02$ ,  $p = 0.13$ ).

### Self-monitoring ability and symptoms (Fig. 3)

In the non-patient groups (i.e. the combined group of healthy controls and the unaffected siblings), detection accuracy over the different conditions was associated with positive schizotypy as measured by the SIS-R ( $\beta = -0.16$ ,  $\text{s.e.} = 0.07$ ,  $p = 0.026$ ). This was not the case for negative schizotypy ( $\beta = -0.05$ ,  $\text{s.e.} = 0.12$ ,  $p = 0.694$ ).

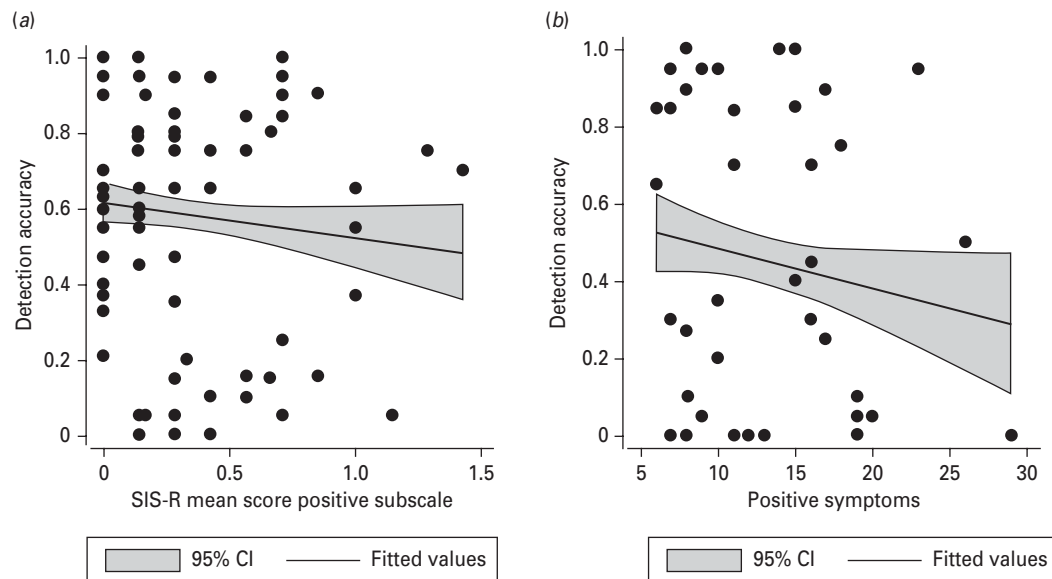
In patients, the level of positive psychotic symptomatology was not robustly associated with detection accuracy over the different conditions ( $\beta = -0.01$ ,  $\text{s.e.} = 0.01$ ,  $p = 0.094$ ). Although the overall three-way condition  $\times$  symptoms  $\times$  antipsychotic use interaction was not statistically significant [interaction  $\chi^2(1) = 1.4$ ,  $p = 0.232$ ], probably because of limited power, further stratification revealed a borderline significant association in patients not using antipsychotic medication ( $\beta = -0.03$ ,  $\text{s.e.} = 0.01$ ,  $p = 0.052$ ), whereas no association was apparent in patients on antipsychotic medication ( $\beta = -0.01$ ,  $\text{s.e.} = 0.01$ ,  $p = 0.407$ ). A similar pattern of associations was found for negative symptoms. There was a suggestion of an association



**Table 2.** Differences in self-monitoring in the five conditions between the genetic risk groups

| Condition | Controls | First-degree relatives |               |       | Patients |                |       |
|-----------|----------|------------------------|---------------|-------|----------|----------------|-------|
|           |          | $\beta$                | 95% CI        | $p$   | $\beta$  | 95% CI         | $p$   |
| 1:1       | N.A.     | 0.02                   | −0.01 to 0.04 | 0.216 | 0.03     | 0.01 to 0.06   | 0.018 |
| 1:2       | N.A.     | 0.01                   | −0.03 to 0.06 | 0.660 | 0.05     | 0.01 to 0.06   | 0.027 |
| 1:4       | N.A.     | −0.08                  | −0.20 to 0.04 | 0.196 | −0.06    | −0.18 to 0.05  | 0.277 |
| 1:6       | N.A.     | −0.15                  | −0.30 to 0.00 | 0.051 | −0.18    | −0.33 to −0.43 | 0.010 |
| 1:8       | N.A.     | −0.09                  | −0.25 to 0.06 | 0.249 | −0.23    | −0.37 to −0.08 | 0.002 |

CI, Confidence interval; N.A., not applicable.

**Fig. 3.** Scatter plots of the detection-symptoms relationship in the most discriminative condition (1:6, see Table 2) in (a) unaffected participants and (b) patients with psychotic disorder. SIS-R, Structured Interview for Schizotypy – Revised; CI, confidence interval.

between negative symptoms and detection accuracy ( $\beta = -0.01$ ,  $\text{s.e.} = 0.01$ ,  $p = 0.064$ ), which was much larger in patients not using antipsychotic medication ( $\beta = -0.04$ ,  $\text{s.e.} = 0.02$ ,  $p = 0.052$ ), and which was absent in patients on antipsychotic treatment ( $\beta = -0.01$ ,  $\text{s.e.} = 0.01$ ,  $p = 0.167$ ), whereas the overall three-way interaction was non-significant [interaction  $\chi^2(1) = 2.2$ ,  $p = 0.135$ ]. Covarying for indicators of severity of illness, that is the number of psychotic episodes and age at first onset, did not alter the results (data not shown).

#### Compensation for mapping change

The distance of the compensation movements increased with the acceleration in all three groups. Multilevel regression analysis revealed no significant differences in compensation among groups over all

mapping changes [group  $\times$  condition interaction:  $\chi^2(2) = 1.66$ ,  $p = 0.44$ ].

#### Discussion

This study assessed the relationship between self-monitoring in healthy controls, patients with psychosis and their unaffected siblings using an action-monitoring task. Several findings emerged. Detection accuracy increased if the mapping change was more obvious in all three groups, but this pattern was more pronounced in the healthy controls than in the patients. Detection accuracy of the unaffected siblings was at an intermediate level between patients and controls. In addition, associations between detection accuracy and symptoms were found. In the combined unaffected groups, an association between detection accuracy and positive schizotypy,

but not negative schizotypy, was shown. In patients, associations between detection accuracy and symptoms were less convincing, although there was suggestive evidence that this association may only be detectable in patients off antipsychotic medication.

The finding that self-monitoring in patients with psychosis was significantly worse than in healthy controls is in line with a large body of work (McGuire *et al.* 1995; Rankin & O'Carroll, 1995; Stirling *et al.* 1998, 2001; Keefe *et al.* 1999, 2002; Bocker *et al.* 2000; Frith *et al.* 2000; Franck *et al.* 2001; Ditman & Kuperberg, 2005; Woodward *et al.* 2007). The finding that unaffected siblings also show worse self-monitoring than healthy controls, but better than their affected brother or sister, is in agreement with two studies that have assessed self-monitoring in groups at increased risk for psychotic disorder (Versmissen *et al.* 2007b; Johns *et al.* 2009). Johns *et al.* (2009) found a significant difference in self-monitoring in persons with an ARMS compared to healthy controls whereas Versmissen *et al.* (2007b) found evidence for self-monitoring alterations in three psychometrically and genetically defined at-risk groups, in a comparison with well controls. These findings may thus suggest that self-monitoring alterations are not merely epiphenomena of acute psychosis in patients. The hypothesis that alterations in the process of the monitoring of one's motor actions may have relevance for the aetiology of psychotic disorder carries the implicit assumption that action-monitoring alterations are associated with psychological processes relevant for the development of psychotic symptoms. Following this reasoning, a global rather than a modality-specific self-monitoring deficit (i.e. limited to the monitoring of one's motor actions) is most likely to increase the risk for psychotic disorder. Future work should address this hypothesis empirically.

In addition, the finding that self-monitoring may be associated with positive symptoms at the level of the disorder and also at the level of subclinical expression suggests that alterations in self-monitoring may have aetiological relevance, as suggested by previous experimental and neuroimaging studies (see Allen *et al.* 2007 for an overview). Nevertheless, the present results do not entirely support previous claims of alterations in self-monitoring as a so-called 'endophenotype' or 'intermediate phenotype' for psychosis (Versmissen *et al.* 2007b), given the possible moderating role of antipsychotic use in the association between self-monitoring and expression of psychosis, which is not in agreement with the notion that endophenotypes are characterized by stability over time and independence of phase of the illness and treatment (Gottesman & Gould, 2003). However, the three-way condition  $\times$  symptoms  $\times$  antipsychotic use interaction was not

significant, because of limited power, indicating that any suggestion of moderation by antipsychotics should be interpreted with caution. If replicated, however, it may be one reason for the non-replication of the association between symptoms and self-monitoring in several previous studies (Vinogradov *et al.* 1997; King, 1998; Fournier *et al.* 2001; Li *et al.* 2002; Moller, 2003; Erhart *et al.* 2006; Versmissen *et al.* 2007a). A similar pattern of results was found for negative symptoms in patients, but this association was not found in the unaffected groups, suggesting that the association with psychotic symptoms may be primary, especially given the known overlap between positive and negative symptoms in patients, which is difficult to account for statistically (Kay, 1990).

Although the detection accuracy of non-self-generated actions was impaired among patients with psychotic disorder and their unaffected siblings, the groups were not impaired in their ability to automatically compensate for the mismatch between self-generated action and their consequences. This was expected and is in accordance with previous work (Kircher & Leube, 2003; Knoblich *et al.* 2004). This finding confirms the literature showing that automatic processes are usually left intact in patients with psychotic disorder, even though conscious cognitive processes are known to be affected (see Jeannerod, 1999 for an overview).

Previous work has claimed that impaired verbal memory, guessing biases or generalized cognitive impairment, rather than alterations in self-monitoring *per se*, may underlie the observation of worse self-monitoring performance in psychosis (Keefe *et al.* 2002). In addition, the use of signal detection tasks using detection of distorted feedback of the patients' speech, which has been applied in previous research (Johns & McGuire, 1999; Johns *et al.* 2001; Fu *et al.* 2006), has also been criticized because these tasks may pick up problems with the appraisal of distorted stimuli instead of problems related to the self-monitoring of the intention to generate (verbal) material (Levitt, 1983). This is supported by the observation of patients with auditory verbal hallucinations and also non-clinical subjects making misattributions of their own voice while passively hearing recorded tapes (Allen *et al.* 2004, 2006). An action-monitoring task, as applied in the present study, is less prone to these possible sources of bias, which is a strength of the study, as is the use of a genetic at-risk group (the unaffected siblings). An important limitation of the study is the small number of patients with psychosis who were not on current antipsychotic treatment. A further limitation is the significant difference in IQ between the three groups tested. Although differences in IQ were present, it is

unlikely that this explains the present results because we controlled statistically for IQ *a priori*, and IQ was only moderately associated with self-monitoring accuracy in the patients but not in the controls or unaffected siblings. To a degree, cognitive alterations such as attention deficits or psychomotor slowing present in families affected with psychotic disorder could explain the observed between-group differences in self-monitoring. Although we did not observe this effect with regard to psychomotor slowing, attention was not specifically assessed in this sample. In addition, environmental factors that are associated with both psychotic disorder and attention deficits, such as cannabis use, could be a source of unmeasured confounding.

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### Declaration of Interest

None.

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